### **CLAIM AMENDMENTS:**

This listing of claims will replace all prior versions and listing of claims in the application. Listing of the Claims:

Claim 1 (currently amended): A quinazoline derivative of the Formula I:

$$R^{1a}$$
 $R^{1b}$ 
 $R^{1b}$ 
 $R^{1b}$ 
 $R^{1b}$ 

wherein:

one of R<sup>1a</sup> or R<sup>1b</sup> is a group of sub-formula (i)

$$Q^2-X^1-Z-Q^1-X^2-O-$$

where  $X^2$  and  $X^1$  are independently selected from a direct bond or a group -[CR<sup>4</sup>R<sup>5</sup>]<sub>m</sub>, wherein m is an integer from 1 to 6,

Z is C(O), SO<sub>2</sub>, -C(O)NR<sup>10</sup>-, -N(R<sup>10</sup>)C(O)-, -C(O)O- or -OC(O)- where R<sup>10</sup> is hydrogen or (1-6C)alkyl,

and each of R<sup>4</sup> and R<sup>5</sup> is independently selected from hydrogen, hydroxy, (1-4C)alkyl, halo(1-4C)alkyl, hydroxy (1-4C)alkyl, (1-4C)alkoxy(1-4C)alkyl, or R<sup>4</sup> and R<sup>5</sup> together with the carbon atom(s) to which they are attached form a (3-7)cycloalkyl ring, provided that when a group R<sup>4</sup> or R<sup>5</sup> is hydroxy, m is at least 2 and the carbon atom to which the hydroxy group is attached is not also attached to another oxygen or a nitrogen atom;

Q<sup>1</sup> is a piperidinyl ring-(3-7C)cycloalkylene or heterocyclyl group, which is optionally substituted by one or two substituents selected from halogeno, trifluoromethyl, trifluoromethoxy,

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cyano, nitro, hydroxy, amino, carboxy, carbamoyl, acryloyl, (1-6C)alkyl, (2-8C)alkenyl,
(2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio,
(2-6C)alkenylthio, (2-6C)alkynylthio, (1-6C)alkylsulfinyl, (2-6C)alkenylsulfinyl,
(2-6C)alkynylsulfinyl, (1-6C)alkylsulfonyl, (2-6C)alkenylsulfonyl, (2-6C)alkynylsulfonyl,
(1-6C)alkylamino, di-[(1-6C)alkylamino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl,
N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino,
N-(1-6C)alkyl-(2-6C)alkanoylamino, sulfamoyl, N-(1-6C)alkylsulfamoyl,
N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino,
N-(1-6C)alkyl-(1-6C)alkanesulfonylamino, carbamoyl(1-6C)alkyl,
N-(1-6C)alkylcarbamoyl(1-6C)alkyl, N,N-di-[(1-6C)alkyl]carbamoyl(1-6C)alkyl,
sulfamoyl(1-6C)alkyl, N-(1-6C)alkylsulfamoyl(1-6C)alkyl,
N,N-di-[(1-6C)alkyl]sulfamoyl(1-6C)alkyl, (2-6C)alkanoyl(1-6C)alkyl,
(2-6C)alkanoyloxy(1-6C)alkyl, (2-6C)alkanoylamino(1-6C)alkyl,
N-(1-6C)alkyl-(2-6C)alkanoylamino(1-6C)alkyl and (1-6C)alkoxycarbonyl(1-6C)alkyl;
Q<sup>2</sup> is an <u>isoxazolyl ring aryl or heteroaryl group</u>, said aryl or heteroaryl group being optionally
substituted by one of more substituents selected from halogeno, trifluoromethyl,
trifluoromethoxy, cyano, nitro, hydroxy, amino, carboxy, carbamoyl, acryloyl, (1-6C)alkyl,
(2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy,
(1-6C)alkylthio, (2-6C)alkenylthio, (2-6C)alkynylthio, (1-6C)alkylsulfinyl,
(2-6C)alkenylsulfinyl, (2-6C)alkynylsulfinyl, (1-6C)alkylsulfonyl, (2-6C)alkenylsulfonyl,
(2-6C)alkynylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl,
\underline{N}-(1-6C)alkylcarbamoyl, \underline{N}, \underline{N}-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy,
(2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, sulfamoyl, N-(1-6C)alkylsulfamoyl,
N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino,
N-(1-6C)alkyl-(1-6C)alkanesulfonylamino, carbamoyl(1-6C)alkyl,
N-(1-6C)alkylcarbamoyl(1-6C)alkyl, N,N-di-[(1-6C)alkyl]carbamoyl(1-6C)alkyl,
sulfamoyl(1-6C)alkyl, N-(1-6C)alkylsulfamoyl(1-6C)alkyl,
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N,N-di-[(1-6C)alkyl]sulfamoyl(1-6C)alkyl, (2-6C)alkanoyl(1-6C)alkyl,

(2-6C)alkanoyloxy(1-6C)alkyl, (2-6C)alkanoylamino(1-6C)alkyl,

N-(1-6C)alkyl-(2-6C)alkanoylamino(1-6C)alkyl and (1-6C)alkoxycarbonyl(1-6C)alkyl, and wherein any (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl and (2-6C)alkanoyl substituent on Q¹ or Q² optionally bears one or more substituents (for example 1, 2 or 3) which may be the same or different selected from halogeno, hydroxy and (1-6C)alkyl and/or optionally a substituent selected from cyano, nitro, carboxy, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, hydroxy(1-6C)alkoxy, (1-4C)alkoxy(1-6C)alkoxy, (2-6C)alkanoyl, (2-6C)alkanoyloxy and NRaRb, wherein Ra is hydrogen or (1-4C)alkyl and Rb is hydrogen or (1-4C)alkyl, and wherein any (1-4C)alkyl in Ra or Rb optionally bears one or more substituents (for example 1, 2 or 3) which may be the same or different selected from halogeno and hydroxy and/or optionally a substituent selected from cyano, nitro, (2-4C)alkenyl, (2-4C)alkynyl, (1-4C)alkoxy, hydroxy(1-4C)alkoxy and (1-2C)alkoxy(1-4C)alkoxy,

or R<sup>a</sup> and R<sup>b</sup> together with the nitrogen atom to which they are attached form a 4, 5 or 6 membered ring, which optionally bears 1 or 2 substituents, which may be the same or different, on an available ring carbon atom selected from halogeno, hydroxy, (1-4C)alkyl and (1-3C)alkylenedioxy, and may optionally bear on any available ring nitrogen a substituent (provided the ring is not thereby quaternised) selected from (1-4C)alkyl, (2-4C)alkanoyl and (1-4C)alkylsulfonyl,

and wherein any (1-4C)alkyl or (2-4C)alkanoyl group present as a substituent on the ring formed by R<sup>a</sup> and R<sup>b</sup> together with the nitrogen atom to which they are attached, optionally bears one or more substituents-(for example 1, 2 or 3) which may be the same or different selected from halogeno and hydroxy and/or optionally a substituent selected from (1-4C)alkyl and (1-4C)alkoxy;

and wherein  $Q^1$  any heterocyclyl-group  $Q^1$ -group optionally bears 1 or 2 oxo (=O) or thioxo (=S) substituents;

and the other of  $R^{1a}$  or  $R^{1b}$  is a group  $R^{1}$  which is selected from hydrogen, hydroxy, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, or a group of the formula:

$$O^4 - X^3 -$$

wherein X³ is a direct bond or is selected from O or S, and Q⁴ is (3-7C)cycloalkyl, (3-7C)cycloalkyl-(1-6C)alkyl, (3-7C)cycloalkenyl-(1-6C)alkyl, heterocyclyl

or heterocyclyl-(1-6C)alkyl,

and wherein adjacent carbon atoms in any (2-6C)alkylene chain within a  $R^1$  substituent are optionally separated by the insertion into the chain of a group selected from O, S, SO, SO<sub>2</sub>, N(R<sup>4</sup>), CO, CH(OR<sup>4</sup>), CON(R<sup>4</sup>), N(R<sup>4</sup>)CO, SO<sub>2</sub>N(R<sup>4</sup>), N(R<sup>4</sup>)SO<sub>2</sub>, CH=CH and C=C wherein R<sup>4</sup> is hydrogen or (1-6C)alkyl,

and wherein any CH<sub>2</sub>=CH- or HC=C- group within a R<sup>1</sup> substituent optionally bears at the terminal CH<sub>2</sub>= or HC= position a substituent selected from halogeno, carboxy, carbamoyl, (1-6C)alkoxycarbonyl,  $\underline{N}$ -(1-6C)alkylcarbamoyl,  $\underline{N}$ - $\underline{N}$ -di-[(1-6C)alkyl]carbamoyl, amino-(1-6C)alkyl, (1-6C)alkylamino-(1-6C)alkyl and di-[(1-6C)alkyl]amino-(1-6C)alkyl or from a group of the formula:

$$0^5 - X^4 -$$

wherein X<sup>4</sup> is a direct bond or is selected from CO and N(R<sup>5</sup>)CO, wherein R<sup>5</sup> is hydrogen or (1-6C)alkyl, and Q<sup>5</sup> is heterocyclyl or heterocyclyl-(1-6C)alkyl,

and wherein any alkyl or alkylene group within a  $R^1$  substituent optionally bears one or more halogeno, (1-6C)alkyl, hydroxy, cyano, amino, carboxy, carbamoyl, sulfamoyl, (1-6C)alkoxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl,  $\underline{N}$ -(1-6C)alkylcarbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino,  $\underline{N}$ -(1-6C)alkyl-(2-6C)alkanoylamino,  $\underline{N}$ -(1-6C)alkylsulfamoyl,  $\underline{N}$ - $\underline{N}$ -di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanosulfonylamino and  $\underline{N}$ -(1-6C)alkyl-(1-6C)alkanosulfonylamino, or from a group of the formula:

$$-X^{5}-Q^{6}$$

wherein  $X^5$  is a direct bond or is selected from O, S, SO, SO<sub>2</sub>, N(R<sup>6</sup>), CO, CH(OR<sup>6</sup>), CON(R<sup>6</sup>), N(R<sup>6</sup>)CO, SO<sub>2</sub>N(R<sup>6</sup>), N(R<sup>6</sup>)SO<sub>2</sub>, C(R<sup>6</sup>)<sub>2</sub>O, C(R<sup>6</sup>)<sub>2</sub>S and C(R<sup>6</sup>)<sub>2</sub>N(R<sup>6</sup>), wherein R<sup>6</sup> is hydrogen or (1-6C)alkyl, and Q<sup>6</sup> is (3-7C)cycloalkyl, (3-7C)cycloalkyl-(1-6C)alkyl, (3-7C)cycloalkenyl-(1-6C)alkyl, heterocyclyl or heterocyclyl-(1-6C)alkyl,

and wherein any heterocyclyl group within a substituent on R<sup>1</sup> optionally bears 1, 2 or 3 substituents, which may be the same or different, selected from halogeno, trifluoromethyl, cyano,

nitro, hydroxy, amino, carboxy, carbamoyl, formyl, mercapto, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, N-(1-6C)alkylsulfamoyl, N-N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkyl-(1-6C)alkylsulfamoyl, N-N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanosulfonylamino, and N-(1-6C)alkyl-(1-6C)alkanosulfonylamino, or from a group of the formula:

$$-X^{6}-R^{7}$$

wherein  $X^6$  is a direct bond or is selected from O, N(R<sup>8</sup>) and C(O), wherein R<sup>8</sup> is hydrogen or (1-6C)alkyl, and R<sup>7</sup> is halogeno-(1-6C)alkyl, hydroxy-(1-6C)alkyl, carboxy-(1-6C)alkyl, (1-6C)alkoxy-(1-6C)alkyl, cyano-(1-6C)alkyl, amino-(1-6C)alkyl, (1-6C)alkylamino-(1-6C)alkyl, di-[(1-6C)alkyl]amino-(1-6C)alkyl, (2-6C)alkanoylamino-(1-6C)alkyl, (1-6C)alkoxycarbonylamino-(1-6C)alkyl, carbamoyl-(1-6C)alkyl, N-(1-6C)alkylcarbamoyl-(1-6C)alkyl, N-(1-6C)alkyl, (2-6C)alkanoyl-(1-6C)alkyl) or (1-6C)alkoxycarbonyl-(1-6C)alkyl,

and wherein any heterocyclyl group within a substituent on R<sup>1</sup> optionally bears 1 or 2 oxo or thioxo substituents;

R<sup>2</sup> is selected from hydrogen and (1-6C)alkyl;

each R<sup>3</sup>, which may be the same or different, is selected from halogeno, cyano, nitro, hydroxy, amino, carboxy, carbamoyl, sulfamoyl, trifluoromethyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkyl, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, N-(1-6C)alkylsulfamoyl, and N,N-di-[(1-6C)alkyl]sulfamoyl a is 1, 2, 3, 4 or 5; 
or a pharmaceutically acceptable salt thereof; subject to the following provisos:

- (i) when Q<sup>2</sup> is aryl, then R<sup>1a</sup> is a group of sub-formula (i) defined above and R<sup>1b</sup> is the group R<sup>1</sup> defined above; and
- (ii) the proviso that the compound of formula I is not one of the following:

N-(3,4-dichlorophenyl)-7-[({4-[(3,5-dimethylisoxazol-4-yl)carbonyl]morpholin-2-

yl}methyl)oxy]-6-(methyloxy)quinazolin-4-amine;

N-(3,4-dichlorophenyl)-7-({[4-(furan-3-ylcarbonyl)morpholin-2-yl]methyl}oxy)-6-(methyloxy)quinazolin-4-amine;

7-[({4-[(2-chloropyridin-3-yl)carbonyl]morpholin 2-yl}methyl)oxy]-N-(3,4-dichlorophenyl)-6-(methyloxy)quinazolin 4-amine; or

7-[({4-[(6-chloropyridin-3-yl)carbonyl]morpholin 2-yl}methyl)oxy]-N-(3,4-dichlorophenyl)-6-(methyloxy)quinazolin-4-amine.

Claim 2 (currently amended): The A-quinazoline derivative according to claim 1 any one of the preceding claims wherein  $X^2$  is a direct bond.

Claim 3 (currently amended): The A-quinazoline derivative according to claim 1-or elaim 2, wherein  $R^{1a}$  is a group of sub-formula (i), and  $R^{1b}$  is a group  $R^{1}$  as defined in claim 1.

Claim 4 (currently amended): The A-quinazoline derivative according to claim 1-or elaim 2, wherein  $R^{1a}$  is a group  $R^{1}$ , and  $R^{1b}$  is a group of sub-formula (i) as defined in claim 1.

Claim 5 (currently amended): The A-quinazoline derivative according to claim 1 any one of the preceding claims, wherein R<sup>1</sup> is selected from hydrogen, hydroxy, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, or a group of the formula:

$$Q^4 - X^3 -$$

wherein X<sup>3</sup> is a direct bond or is O or S (particularly a direct bond or O), and Q<sup>4</sup> is (3-7C)cycloalkyl, (3-7C)cycloalkyl-(1-6C)alkyl, (3-7C)cycloalkenyl, (3-7C)cycloalkenyl-(1-6C)alkyl, heterocyclyl or heterocyclyl-(1-6C)alkyl, and wherein any alkyl or alkylene group within a R<sup>1</sup> substituent optionally bears one or more

halogeno, (1-6C)alkyl, hydroxy, cyano, amino, carboxy, carbamoyl, sulfamoyl, (1-6C)alkoxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N-(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, N-(1-6C)alkylsulfamoyl, N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanosulfonylamino and N-(1-6C)alkyl-(1-6C)alkanosulfonylamino.

Claim 6 (currently amended): The A-quinazoline derivative according to claim 5 wherein R<sup>1</sup> is hydrogen, (1-6C)alkoxy and (1-4C)alkoxy(1-6C)alkoxy, and wherein any (1-6C)alkoxy group within R<sup>1</sup> optionally bears 1, 2 or 3 substituents, which may be the same or different, selected from hydroxy, fluoro and chloro.

Claim 7 (**currently amended**): <u>The A-quinazoline derivative according to claim 6</u> wherein R<sup>1</sup> is selected from methoxy, ethoxy, isopropyloxy, cyclopropylmethoxy, 2-hydroxyethoxy, 2-fluoroethoxy, 2-methoxyethoxy, 2,2-difluoroethoxy, 2,2,2-trifluoroethoxy or 3-hydroxy-3-methylbutoxy.

Claim 8 (currently amended): <u>The A-quinazoline derivative according to claim 5</u> wherein R<sup>1</sup> is methoxy.

Claim 9 (currently amended): The A-quinazoline derivative according to claim 1-any one of the preceding claims wherein  $X^1$  is suitably a direct bond or a (1-6C)alkylene group.

Claim 10 (currently amended):  $\underline{\text{The }}$  A-quinazoline derivative according to claim 9 wherein  $X^1$  is a direct bond or methylene or ethylene group.

Claim 11 (currently amended): <u>The A-quinazoline derivative according to claim 1-any</u> one of the preceding claims wherein Z is selected from -C(O)-,  $-NR^{10}-C(O)$ - (wherein  $R^{10}$  is H or (1-6C)alkyl), and -O-C(O)-.

Claim 12 (currently amended): <u>The A-quinazoline derivative according to claim 11</u>, wherein Z is -C(O)-.

Claim 13 (currently amended): <u>The A-quinazoline derivative according to claim 11</u>, wherein Z is selected from -NH-C(O)- and -O-C(O)-.

Claims 14-15 (cancelled).

Claim 16 (currently amended): The A-quinazoline derivative according to claim 11 any one of claims 11 to 16, wherein the group Q<sup>2</sup>-X<sup>1</sup>-Z- is linked to the piperidinyl nitrogen-a nitrogen atom on a heterocyclic atom of Q<sup>1</sup>.

Claim 17 (currently amended): The A-quinazoline derivative according to claim 1-any one of the preceding claims, wherein Q² is a heteroaryl group, said heteroaryl group being optionally substituted by one of more substituents selected from halogeno, trifluoromethyl, trifluoromethoxy, cyano, nitro, hydroxy, amino, carboxy, carbamoyl, acryloyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (2-6C)alkynylthio, (1-6C)alkylsulfinyl, (2-6C)alkynylsulfinyl, (2-6C)alkynylsulfinyl, (1-6C)alkylsulfonyl, (2-6C)alkenylsulfonyl, (2-6C)alkynylsulfonyl, (1-6C)alkylljamino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylsulfonyl, (1-6C)alkylljcarbamoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, sulfamoyl, N-(1-6C)alkylsulfamoyl, N-(1-6C)alkylsulfamoyl, (1-6C)alkylsulfamoyl, N-(1-6C)alkylsulfamoyl, N-(1-6C)alkylsulfamoyl, N-(1-6C)alkylsulfamoyl(1-6C)alkyl, (2-6C)alkanoyl(1-6C)alkyl, N-(1-6C)alkylsulfamoyl(1-6C)alkyl, (2-6C)alkanoyl(1-6C)alkyl, N-(1-6C)alkylsulfamoyl(1-6C)alkyl, (2-6C)alkanoyl(1-6C)alkyl,

(2-6C)alkanoyloxy(1-6C)alkyl, (2-6C)alkanoylamino(1-6C)alkyl,

N-(1-6C)alkyl-(2-6C)alkanoylamino(1-6C)alkyl and (1-6C)alkoxycarbonyl(1-6C)alkyl,
and wherein any (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl and (2-6C)alkanoyl
substituent on Q² optionally bears one or more substituents-(for example 1, 2 or 3) which may be
the same or different selected from halogeno, hydroxy and (1-6C)alkyl and/or optionally a
substituent selected from cyano, nitro, carboxy, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy,
hydroxy(1-6C)alkoxy, (1-4C)alkoxy(1-6C)alkoxy, (2-6C)alkanoyl, (2-6C)alkanoyloxy and
NRaRb, wherein Ra is hydrogen or (1-4C)alkyl and Rb is hydrogen or (1-4C)alkyl, and wherein
any (1-4C)alkyl in Ra or Rb optionally bears one or more substituents (for example 1, 2 or 3)
which may be the same or different selected from halogeno and hydroxy and/or optionally a
substituent selected from cyano, nitro, (2-4C)alkenyl, (2-4C)alkynyl, (1-4C)alkoxy,

or R<sup>a</sup> and R<sup>b</sup> together with the nitrogen atom to which they are attached form a 4, 5 or 6 membered ring, which optionally bears 1 or 2 substituents, which may be the same or different, on an available ring carbon atom selected from halogeno, hydroxy, (1-4C)alkyl and (1-3C)alkylenedioxy, and may optionally bear on any available ring nitrogen a substituent (provided the ring is not thereby quaternised) selected from (1-4C)alkyl, (2-4C)alkanoyl and (1-4C)alkylsulfonyl,

and wherein any (1-4C)alkyl or (2-4C)alkanoyl group present as a substituent on the ring formed by R<sup>a</sup> and R<sup>b</sup> together with the nitrogen atom to which they are attached, optionally bears one or more substituents (for example 1, 2 or 3) which may be the same or different selected from halogeno and hydroxy and/or optionally a substituent selected from (1-4C)alkyl and (1-4C)alkoxy.

Claims 18-22 (cancelled).

hydroxy(1-4C)alkoxy and (1-2C)alkoxy(1-4C)alkoxy,

Claim 23 (currently amended): The A-quinazoline derivative according to claim 1-any one of the preceding claims wherein Q<sup>2</sup> optionally bears 1 or 2 substituents, which may be the same or different, selected from halogeno, hydroxy, nitro, amino, cyano, carbamoyl, (1-4C)alkyl,

(1-4C)alkoxy, (2-4C)alkanoyl and (1-4C)alkylsulfonyl, (1-4C)alkylamino, di[(1-4C)alkyl]amino, *N*-[(1-4C)alkyl]carbamoyl, and *N*,*N*-di[(1-4C)alkyl]carbamoyl.

and wherein any (1-4C)alkyl, or (2-4C)alkanoyl group within Q<sup>2</sup> optionally bears 1 or 2 substituents, which may be the same or different, selected from halogeno, hydroxy and (1-6C)alkyl and/or optionally a substituent selected from cyano, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkanoyl, (2-6C)alkanoyloxy and NR<sup>a</sup>R<sup>b</sup>, wherein R<sup>a</sup> is hydrogen or (1-4C)alkyl and R<sup>b</sup> is hydrogen or (1-4C)alkyl, and wherein any (1-4C)alkyl in R<sup>a</sup> or R<sup>b</sup> optionally bears one or more substituents (for example 1, 2 or 3) which may be the same or different selected from halogeno and hydroxy and/or optionally a substituent selected from cyano, and (1-4C)alkoxy,

or R<sup>a</sup> and R<sup>b</sup> together with the nitrogen atom to which they are attached form a 4, 5 or 6 membered ring which does not contain oxygen, which ring optionally bears 1 or 2 substituents, which may be the same or different, on an available ring carbon atom selected from halogeno, hydroxy, (1-4C)alkyl and (1-3C)alkylenedioxy, and may optionally bear on any available ring nitrogen a substituent (provided the ring is not thereby quaternised) selected from (1-4C)alkyl, (2-4C)alkanoyl and (1-4C)alkylsulfonyl,

and wherein any (1-4C)alkyl or (2-4C)alkanoyl group present as a substituent on the ring formed by R<sup>a</sup> and R<sup>b</sup> together with the nitrogen atom to which they are attached optionally bears one or more substituents (for example 1, 2 or 3), which may be the same or different, selected from halogeno and hydroxy and/or optionally a substituent selected from (1-4C)alkyl and (1-4C)alkoxy.

Claim 24 (**currently amended**): <u>The A-quinazoline derivative according to claim 23</u> wherein  $Q^2$  is optionally substituted by one or two groups, which may be the same or different, selected from halogeno, hydroxy, nitro, amino, cyano, carbamoyl, (1-4C)alkyl, (1-4C)alkoxy, (2-4C)alkanoyl and (1-4C)alkylsulfonyl, [(1-4C)alkyl]amino, di[(1-4C)alkyl]amino, N-[(1-4C)alkyl]carbamoyl, and N-di[(1-4C)alkyl]carbamoyl.

and wherein any (2-4C)alkanoyl group in a substituent on Q<sup>2</sup> optionally bears one or two substituents, which may be the same or different, selected from hydroxy and (1-3C)alkyl,

and wherein any (1-4C)alkyl group in a substituent on Q<sup>2</sup> optionally bears one or two substituents, which may be the same or different, selected from hydroxy, (1-4C)alkoxy and halogeno (particularly chloro and more particularly fluoro).

Claim 25 (**currently amended**): <u>The A-quinazoline derivative according to claim 23-or elaim 24</u> wherein Q<sup>2</sup> is unsubstituted or substituted by a (1-4C)alkyl group, a (1-4C)alkoxy group, halogeno, amino, nitro, cyano, carbamoyl, di-[(1-4C)alkyl]amino, and *N,N*-di[(1-4C)alkyl]carbamoyl.

Claim 26 (currently amended): <u>The A-quinazoline derivative according to claim 1 any</u> one of the preceding claims wherein R<sup>2</sup> is hydrogen.

Claim 27 (currently amended): <u>The A-quinazoline derivative according to claim 1 any</u> one of the preceding claims wherein a is 1, 2 or 3.

Claim 28 (currently amended): The A-quinazoline derivative according to claim 1-any one of the preceding claims, wherein an R<sup>3</sup> is in the para position on the anilino ring, and this is selected from halogeno, cyano, nitro, hydroxy, amino, trifluoromethyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylamino and di-[(1-6C)alkyl]amino.

Claim 29 (currently amended): <u>The A-quinazoline derivative according to claim 1-any</u> one of the preceding claims wherein the group of sub-formula (ii)

in formula (I) is a group of sub-formula (iii)

where one of R<sup>15</sup> or R<sup>17</sup> is hydrogen and the other is halogeno, and R<sup>16</sup> is halogeno.

Claim 30 (currently amended): <u>The A-quinazoline derivative according to claim 29</u> wherein the group of sub-formula (iii) is 3-chloro-2-fluorophenyl, or 3-chloro-4-fluorophenyl.

Claim 31 (currently amended): <u>The A-compound according to claim 1</u> selected from one of the following:

- (1) *N*-(3-chloro-2-fluorophenyl)-6-{[1-(isoxazol-5-ylcarbonyl)piperidin-4-yl]oxy}-7-methoxyquinazolin-4-amine;
- (2) N-(3-chloro-2-fluorophenyl)-7-methoxy-6-({1-[(3-methylisoxazol-5-yl)acetyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (3) *N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-({1-[(3-methylisoxazol-5-yl)carbonyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (4) *N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-({1-[(5-methylisoxazol-3-yl)carbonyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (5) *N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-({1-[(5-methylisoxazol-4-yl)carbonyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (6) N-(3-chloro-2-fluorophenyl)-7-methoxy-6-({1-[(3-methylisoxazol-4-yl)carbonyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (7) *N*-(3-chloro-2-fluorophenyl)-6-({1-[(3,5-dimethylisoxazol-4-yl)carbonyl]piperidin-4-yl}oxy)-7-methoxyquinazolin-4-amine;
- (8)N (3 chloro-2-fluorophenyl)-7-methoxy-6 {[1-(pyridin-3-ylcarbonyl)piperidin-4-yl]oxy}quinazolin-4-amine:

- (9) N (3-chloro-2-fluorophenyl) 7 methoxy-6-{[1-(pyridin-2-ylcarbonyl)piperidin-4-yl]oxy}quinazolin-4-amine;
- (10)N-(3-chloro-2-fluorophenyl)-6-{[1-(2-furoyl)piperidin-4-yl]oxy}-7-methoxyquinazolin-4-amine;
- (11)(8) N-(3-chloro-2-fluorophenyl)-7-{[1-(isoxazol-5-ylcarbonyl)piperidin-4-yl]oxy}-6-methoxyquinazolin-4-amine;
- (12)(9) N-(3-chloro-2-fluorophenyl)-6-methoxy-7-({1-[(3-methylisoxazol-5-yl)acetyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (13)N (3 chloro-2-fluorophenyl) 7-{[1 (pyridin-3-ylcarbonyl)piperidin-4-yl]oxy}-6-methoxyquinazolin-4-amine;
- (14)N (3-chloro 2-fluorophenyl) 7-{[1-(2-furoyl)piperidin-4-yl]oxy}-6-methoxyquinazolin-4-amine;
- (15)N-(3-chloro-2-fluorophenyl)-7-methoxy-6-{[(3R)-1-(2-thienylacetyl)piperidin-3-yl]oxy}quinazolin-4-amine;
- (16)N-(3 chloro 2-fluorophenyl) 6-{[(3R) 1-isonicotinoylpiperidin-3-yl]oxy}-7-methoxyquinazolin-4-amine;
- (17)6-({(3R) 1-[(2-aminopyridin-3-yl)carbonyl]piperidin-3-yl}oxy)-N-(3-chloro-2-fluorophenyl)-7-methoxyquinazolin-4-amine;
- (18)N (3 chloro 2-fluorophenyl) 7-methoxy 6-{[(3R) 1-(1H-pyrrol-2-ylcarbonyl)piperidin 3-yl]oxy}quinazolin-4-amine;
- (19)N-(3-chloro-2-fluorophenyl)-7-methoxy-6- $\{[(3R)-1-(2$ -thienylcarbonyl)piperidin-3-yl]oxy}quinazolin-4-amine;
- (20)N-(3-chloro-2-fluorophenyl)-6-{[(3R)-1-(2-furoyl)piperidin-3-yl]oxy}-7-methoxyquinazolin-4-amine;
- (21)N-(3 chloro-2-fluorophenyl)-6-{[(3R)-1-(3-furoyl)piperidin-3-yl]oxy}-7-methoxyquinazolin-4-amine;
- (22)N-(3-chloro-2-fluorophenyl) 7-methoxy-6-{[(3R)-1-(3-thienylcarbonyl)piperidin-3-yl]oxy}quinazolin 4-amine;

- (23)N (3-chloro-2-fluorophenyl)-7-methoxy-6-{[(3R)-1-(3-thienylacetyl)piperidin-3-yl]oxy}quinazolin-4-amine;
- (24)N-(3-chloro-2-fluorophenyl)-7-methoxy-6-({(3R)-1-[(1-methyl-1H-pyrrol-2-yl)carbonyl]piperidin-3-yl}oxy)quinazolin-4-amine;
- (25)N-(3-chloro-2-fluorophenyl)-7-methoxy-6-({(3R)-1-[(4-nitro-1*H*-pyrazol-1-yl)acetyl]piperidin-3-yl}oxy)quinazolin-4-amine;
- (26)(10) N-(3-chloro-2-fluorophenyl)-7-methoxy-6-((3R)-1-[(3-methylisoxazol-5-yl)acetyl]piperidin-3-yl}oxy)quinazolin-4-amine;
- (27)(11) N-(3-chloro-2-fluorophenyl)-7-methoxy-6-{[(3R)-1-(4-{N,N-dimethylcarbamoyl}-1H-pyrazol-1-ylacetyl)piperidin-3-yl]oxy}quinazolin-4-amine; and
- (28)N (3 chloro-2-fluorophenyl)-7-methoxy-6-{[(3R)-1-(4-cyano-1H-pyrazol-1-ylacetyl)piperidin-3-yl]oxy}quinazolin-4-amine;
- (29)4 ({4-[(3-Chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-phenylpiperidine-1-carboxamide;
- (30)N-Benzyl-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)piperidine-1-carboxamide;
- (31)4-({4-[(3-Chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-[4-(dimethylamino)phenyl]piperidine-1-carboxamide:
- (32)4 ({4-[(3-Chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy) N (2-phenylethyl)piperidine-1-carboxamide;
- (33)4-({4-[(3-Chloro 2 fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy) N (3,4-dimethoxyphenyl)piperidine-1-carboxamide;
- (34)4 ({4-[(3-Chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy) N (3-fluorophenyl)piperidine-1-carboxamide;
- (35)(12) 4-({4-[(3-Chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-(3,5-dimethylisoxazol-4-yl)piperidine-1-carboxamide.
- (36)4-({4-[(3-Chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}-oxy)-N-2-thienylpiperidine-1-carboxamide;

(37)4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-3-thienylpiperidine-1-carboxamide.

Claim 32 (currently amended): A process for the preparation of a quinazoline derivative of the Formula I as defined in any one of the preceding claims, which process comprises either **Process (a)** reacting a compound of the Formula II:

$$R^{1a'}$$
 $R^{1b'}$ 
 $R^{1b'}$ 
 $R^{1b'}$ 

#### Formula II

wherein  $R^3$  and a are as defined in claim 1 and one of  $R^{1a'}$  or  $R^{1b'}$  is hydroxy and the other is a group  $R^1$  as defined in claim 1 in relation to formula (I), except that any functional group is protected if necessary,

with a compound of the Formula III:

$$Q^2-X^1-Z-Q^1-X^2-Lg$$

# Formula III

wherein  $Q^1$ ,  $Q^2$ , Z,  $X^2$  and  $X^1$  have any of the meanings defined in claim 1, except that any functional group is protected if necessary and Lg is a displaceable group:

**Process (b)** modifying a substituent in or introducing a substituent into another quinazoline derivative of Formula I or a pharmaceutically acceptable salt thereof as defined in claim 1, except that any functional group is protected if necessary;

**Process (c)** reacting a compound of the Formula II as defined in respect of process (a) above with a compound of the Formula III as defined in process (a) except Lg is OH under Mitsunobu conditions,

**Process (d)** for the preparation of those compounds of the Formula I wherein the group R<sup>1</sup> is a hydroxy group by the cleavage of a quinazoline derivative of the Formula I wherein R<sup>1</sup> is a (1-6C)alkoxy group;

**Process (e)** for the preparation of those compounds of the Formula I wherein R<sup>1</sup> is a (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, or a group of the formula:

$$O^4 - X^3 -$$

wherein  $X^3$  is O and  $Q^4$  is as defined in claim 5, by the reaction of a compound of the Formula I wherein  $R^1$  is OH, except that any functional group is protected if necessary, with a compound of the formula  $R^{1'}$ -Lg, wherein  $R^{1'}$  is a (1-6C)alkyl, (2-6C)alkenyl, (2-6C)alkynyl, or a group  $Q^4$  where  $Q^4$  is as defined in claim 5, and Lg is a displaceable group;

**Process (f)** for the preparation of those compounds of the Formula I wherein  $Q^1$ ,  $Q^2$  contains or  $R^1$  is or contains a (1-6C)alkoxy or substituted (1-6C)alkoxy group or a (1-6C)alkylamino or substituted (1-6C)alkylamino group, the alkylation of a quinazoline derivative of the Formula I wherein  $Q^1$ ,  $Q^2$  contains or  $R^1$  is or contains a hydroxy group or a primary or secondary amino group as appropriate;

**Process (g)** for the preparation of those compounds of the Formula I wherein  $R^1$  is substituted by a group T, wherein T is selected from (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (2-6C)alkanoylamino, (1-6C)alkylthio, (1-6C)alkylsulfinyl and (1-6C)alkylsulfonyl, the reaction of a compound which is of formula (I) except that the group  $R^1$  is replaced with a group  $R^1$ "-Lg wherein Lg is a displaceable group, and  $R^1$ " is a group  $R^1$  except that it has Lg in place of the group T, and further that any functional group is protected if necessary, with a compound of the formula TH, wherein T is as defined above except that any functional group is protected if necessary;

Process (h) by reacting a compound of the formula VI:

#### formula VI

wherein R<sup>1a</sup> and R<sup>1b</sup> have any of the meanings defined in claim 1 except that any functional group is protected if necessary and Lg is a displaceable group, with an aniline of the formula VII:

# formula VII

wherein R<sup>3</sup> and a have any of the meanings defined in claim 1, except that any functional group is protected if necessary, and wherein the reaction is conveniently performed in the presence of a suitable acid, or

**Process (i)** for the preparation of those compounds of the Formula I wherein  $Q^1$  is a nitrogen containing heterocyclyl group linked to the group Z by a ring nitrogen, the coupling of a compound of the Formula I as defined in claim 1, except that the group of sub-formula (i) is a group of sub-formula (x) H- $Q^1$ - $X^2$ -O-, and any functional group is protected if necessary, with a compound of formula  $Q^2$ - $X^1$ -Z-Lg, wherein Z,  $Q^2$  and  $X^1$  are as defined in claim 1 and Lg is a leaving group;

**Process (j)** for the preparation of those compounds of the Formula I define in claim 1 wherein  $Q^1$  is a nitrogen containing heterocyclyl group linked to the -Z- group by a ring nitrogen, and Z is a group of formula -NR<sup>10</sup>-C(O)-; said process comprising the coupling of a compound of the Formula I, except that the group of sub-formula (i) is a group of sub-formula (x) H-Q<sup>1</sup>-X<sup>2</sup>-O-, and any functional group is protected if necessary, with a compound of formula Q<sup>2</sup>-X<sup>1</sup>-N=C=O, wherein Q<sup>2</sup> and X<sup>1</sup> are as defined in claim 1;

and whereafter any protecting group that is present is removed-by conventional means.

Claim 33 (currently amended): <u>The A-process</u> according to claim 32, wherein Lg is a leaving group selected from hydroxyl, chloro or bromo.

Claim 34 (currently amended): A pharmaceutical composition which comprises a quinazoline derivative of the Formula I, or a pharmaceutically-acceptable salt thereof, as defined in claim 1 any one of claims 1 to 31 in association with a pharmaceutically-acceptable diluent or carrier.

Claims 35-36 (cancelled).

Claim 37 (**currently amended**): A method for producing an anti-proliferative effect in a warm-blooded animal in need of such treatment which comprises administering to said animal a quinazoline derivative of the Formula I, or a pharmaceutically acceptable salt thereof, as defined in <u>claim 1</u> any one of claims 1 to 31.